

1

SEQUENCE LISTING

<110> Kindsvogel, Wayne R. Topouzis, Stavros

| <120> SOLUBLE ZCYTOR11 CYTOKINE RECEPTORS | | | | | | | | | | | | | | |
|---|--|--|--|--|--|--|--|--|--|--|--|--|--|--|
| <130> 00-56 | | | | | | | | | | | | | | |
| <150> US 60/223,827 <151> 2000-08-08 | | | | | | | | | | | | | | |
| <150> US 60/250,876 <151> 2000-12-01 | | | | | | | | | | | | | | |
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| ttg act gtg gga tcc ctg gct gct cac gcc cct gag gac ccc tcg gat 102 Leu Thr Val Gly Ser Leu Ala Ala His Ala Pro Glu Asp Pro Ser Asp 10 15 20 | | | | | | | | | | | | | | |
| ctg ctc cag cac gtg aaa ttc cag tcc agc aac ttt gaa aac atc ctg 150 Leu Leu Gln His Val Lys Phe Gln Ser Ser Asn Phe Glu Asn Ile Leu 25 30 35 | | | | | | | | | | | | | | |
| acg tgg gac agc ggg cca gag ggc acc cca gac acg gtc tac agc atc 198 Thr Trp Asp Ser Gly Pro Glu Gly Thr Pro Asp Thr Val Tyr Ser Ile 40 45 50 55 | | | | | | | | | | | | | | |
| gag tat aag acg tac gga gag agg gac tgg gtg gca aag aag ggc tgt 246 Glu Tyr Lys Thr Tyr Gly Glu Arg Asp Trp Val Ala Lys Lys Gly Cys 60 65 70 | | | | | | | | | | | | | | |
| cag cgg atc acc cgg aag tcc tgc aac ctg acg gtg gag acg ggc aac 294 Gln Arg Ile Thr Arg Lys Ser Cys Asn Leu Thr Val Glu Thr Gly Asn 75 80 85 | | | | | | | | | | | | | | |
| ctc acg gag ctc tac tat gcc agg gtc acc gct gtc agt gcg gga ggc 342 Leu Thr Glu Leu Tyr Tyr Ala Arg Val Thr Ala Val Ser Ala Gly Gly 90 95 100 | | | | | | | | | | | | | | |
| cgg tca gcc acc aag atg act gac agg ttc agc tct ctg cag cac act 390 Arg Ser Ala Thr Lys Met Thr Asp Arg Phe Ser Ser Leu Gln His Thr 105 110 | | | | | | | | | | | | | | |
| acc ctc aag cca cct gat gtg acc tgt atc tcc aaa gtg aga tcg att | | | | | | | | | | | | | | |

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|-------------------|------------|-------------------|-------------------|------------|-------------------|------------|-------------------|-------------------|------------|-------------------|------------|-------------------|-------------------|------------|-------------------|------|
| | | | | | | | | acg Thr | | | | | | | | 486 |
| | | | | | | | | ttc Phe 160 | | | | | | | | 534 |
| gag Glu | ctc Leu | cag Gln 170 | gtc Val | aac Asn | cgc Arg | acc Thr | tac Tyr 175 | caa Gln | atg Met | cac His | ctt Leu | gga Gly 180 | ggg Gly | aag Lys | cag Gln | 582 |
| | | | | | | | | acc Thr | | | | | | | | 630 |
| | | | | | | | | tgg Trp | | | | | | | | 678 |
| | | | | | | | | gac Asp | | | | | | | | 726 |
| tcc Ser | gga Gly | gcc Ala | ttc Phe 235 | ctg Leu | ttc Phe | tcc Ser | atg Met | ggc Gly 240 | ttc Phe | ctc Leu | gtc Val | gca Ala | gta Val 245 | ctc Leu | tgc Cys | 774 |
| | | | | | | | | aag Lys | | | | | | | | 822 |
| | | | | | | | | ttc Phe | | | | | | | | 870 |
| | | | | | | | | gac Asp | | | | | | | | 918 |
| | | | | | | | | atc Ile | | | | | | | | 966 |
| | | | | | Gln | Arg | His | agc Ser 320 | Leu | Ser | Glu | | | | | 1014 |
| | | | | | | | | cag Gln | | | | | | | | 1062 |
| | | | | | | | | gcc Ala | | | | | | | | 1110 |
| ggg Gly 360 | ccc Pro | cca Pro | tcc Ser | tat Tyr | gca Ala 365 | cct Pro | cag Gln | gtg Val | acc Thr | ccc Pro 370 | gaa Glu | gct Ala | caa Gln | ttc Phe | cca Pro 375 | 1158 |
| | | | | | | | | aag Lys | | | | | | | | 1206 |
| | | | | | | | | cct Pro 400 | | | | | | | | 1254 |
| gaa | ggt | tct | ggc | aaa | gac | tcc | ccc | act | ggg | aca | ctt | tct | agt | cct | aaa | 1302 |

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| Glu | Gly | Ser 410 | Gly | Lys | Asp | Ser | Pro 415 | Thr | Gly | 3 Thr | Leu | ser 420 | ser | Pro | Lys | |
|--|---|---|---|---|--|--|--|--|---|--|---|--|---|---|--|---|
| cac His | ctt Leu 425 | agg Arg | cct Pro | aaa Lys | ggt Gly | cag Gln 430 | ctt Leu | cag Gln | aaa Lys | gag Glu | cca Pro 435 | cca Pro | gct Ala | gga Gly | agc Ser | 1350 |
| tgc Cys 440 | atg Met | tta Leu | ggt Gly | ggc Gly | ctt Leu 445 | tct Ser | ctg Leu | cag Gln | gag Glu | gtg Val 450 | acc Thr | tcc Ser | ttg Leu | gct Ala | atg Met 455 | 1398 |
| gag Glu | gaa Glu | tcc Ser | caa Gln | gaa Glu 460 | gca Ala | aaa Lys | tca Ser | ttg Leu | cac His 465 | cag Gln | ccc Pro | ctg Leu | ggg Gly | att Ile 470 | tgc Cys | 1446 |
| aca Thr | gac Asp | aga Arg | aca Thr 475 | tct Ser | gac Asp | cca Pro | aat Asn | gtg Val 480 | cta Leu | cac His | agt Ser | ggg Gly | gag Glu 485 | gaa Glu | ggg Gly | 1494 |
| aca Thr | cca Pro | cag Gln 490 | tac Tyr | cta Leu | aag Lys | ggc Gly | cag Gln 495 | ctc Leu | ccc Pro | ctc Leu | ctc Leu | tcc ser 500 | tca Ser | gtc Val | cag Gln | 1542 |
| | | | cac His | | | | | | | | | | | | | 1590 |
| tgt Cys 520 | tcc Ser | ccc Pro | tcg Ser | gac Asp | caa Gln 525 | ggt Gly | cca Pro | agt Ser | ccc Pro | tgg Trp 530 | ggc Gly | ctg Leu | ctg Leu | gag Glu | tcc Ser 535 | 1638 |
| ctt Leu | gtg Val | tgt Cys | ccc Pro | aag Lys 540 | gat Asp | gaa Glu | gcc Ala | aag Lys | agc ser 545 | cca Pro | gcc Ala | cct Pro | gag Glu | acc Thr 550 | tca Ser | 1686 |
| gac Asp | ctg Leu | gag Glu | cag Gln 555 | ccc Pro | aca Thr | gaa Glu | ctg Leu | gat Asp 560 | tct Ser | ctt Leu | ttc Phe | aga Arg | ggc Gly 565 | ctg Leu | gcc Ala | 1734 |
| | | | cag Gln | | | | tga | gggg | aat (| gggaa | aagg | ct t | ggtg | cttc | C | 1785 |
| ggcgggaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaaa | totgo cocto tgaca cocato gacaa acaco tctgo gtggo ccaaa o 2 2 1> 52 PI | cga gctgctgggaacctggcctggaaccggaaccggaacctgg | tctggatgggagggagggaggggaggggggggggggggg | gccte gtgcg ggage gaage gaaage ttcate aacgte aacgge accate acgge | ca go gc to gct a to gccggaac co ggaac co ggaac ggaac ggaac co ggaac co ggaac co ggaac co ggaac co ggaac co gga | acgggcctcagtgcaggcctgaaaggcctgaaaggcctgaaaggcctgaaaggcctgaaaggcctggcccagtggcccagtggccagtgccagt | gtgcoggacacacacacacacacacacacacacacacacaca | c cttg aad agg agg gg g | tgaga caaa cctcc cctc tgcc aaga cagca tcag tcag | agaa gcag tgct cgag tcctt acca agga cctga cctag attga ctta | gcaç catç gtc tgc ctg gaa cgg acg ggt tca acc | gaggggataggataggaggagggagggaggaggaggagga | gag agg ccc gtt gtt gct acg acg acg acc acc acc acca acca | tggca actg tgca ccac cctgg tcaa gggc actt agct agc | gccaca atgcag aggcag ctccagtt aggtgg agaacc ggaggctt aagcctg gtgagcta gtggatca agtgcta | 1905 1965 2025 2085 2145 2205 2325 2325 2345 2505 265 265 265 265 265 2745 |
| | | | | | | | | | | | | | | | | |

 $<\!400\!>\,2$ Met Arg Thr Leu Leu Thr Ile Leu Thr Val Gly Ser Leu Ala Ala His

5 10 Ala Pro Glu Asp Pro Ser Asp Leu Leu Gln His Val Lys Phe Gln Ser 20 25 30 Ser Asn Phe Glu Asn Ile Leu Thr Trp Asp Ser Gly Pro Glu Gly Thr 40 Pro Asp Thr Val Tyr Ser Ile Glu Tyr Lys Thr Tyr Gly Glu Arg Asp 50 55 60 Trp Val Ala Lys Lys Gly Cys Gln Arg Ile Thr Arg Lys Ser Cys Asn 65 70 75 80 Leu Thr Val Glu Thr Gly Asn Leu Thr Glu Leu Tyr Tyr Ala Arg Val Thr Ala Val Ser Ala Gly Gly Arg Ser Ala Thr Lys Met Thr Asp Arg 105 Phe Ser Ser Leu Gln His Thr Thr Leu Lys Pro Pro Asp Val Thr Cys
115 120 125 Ile Ser Lys Val Arg Ser Ile Gln Met Ile Val His Pro Thr Pro Thr 135 Pro Île Arg Ala Gly Asp Gly His Arg Leu Thr Leu Glu Asp Île Phe 145 150 155 160 His Asp Leu Phe Tyr His Leu Glu Leu Gln Val Asn Arg Thr Tyr Gln
165 170 175 Met His Leu Gly Gly Lys Gln Arg Glu Tyr Glu Phe Phe Gly Leu Thr 180 185 190 185 Pro Asp Thr Glu Phe Leu Gly Thr Ile Met Ile Cys Val Pro Thr Trp 200 Ala Lys Glu Ser Ala Pro Tyr Met Cys Arg Val Lys Thr Leu Pro Asp 210 215 220 Arg Thr Trp Thr Tyr Ser Phe Ser Gly Ala Phe Leu Phe Ser Met Gly 225 230 Phe Leu Val Ala Val Leu Cys Tyr Leu Ser Tyr Arg Tyr Val Thr Lys 245 250 255 Pro Pro Ala Pro Pro Asn Ser Leu Asn Val Gln Arg Val Leu Thr Phe 265 260 Gln Pro Leu Arg Phe Ile Gln Glu His Val Leu Ile Pro Val Phe Asp 280 Leu Ser Gly Pro Ser Ser Leu Ala Gln Pro Val Gln Tyr Ser Gln Ile 295 300 Arg Val Ser Gly Pro Arg Glu Pro Ala Gly Ala Pro Gln Arg His Ser 31Ŏ 315 Leu Ser Glu Ile Thr Tyr Leu Gly Gln Pro Asp Ile Ser Ile Leu Gln 325 330 Pro Ser Asn Val Pro Pro Pro Gln Ile Leu Ser Pro Leu Ser Tyr Ala 340 345 350 345 340 Pro Asn Ala Ala Pro Glu Val Gly Pro Pro Ser Tyr Ala Pro Gln Val 360 365 Thr Pro Glu Ala Gln Phe Pro Phe Tyr Ala Pro Gln Ala Ile Ser Lys 375 Val Gln Pro Ser Ser Tyr Ala Pro Gln Ala Thr Pro Asp Ser Trp Pro 385 390 395 400 Pro Ser Tyr Gly Val Cys Met Glu Gly Ser Gly Lys Asp Ser Pro Thr 405 410 415 Gly Thr Leu Ser Ser Pro Lys His Leu Arg Pro Lys Gly Gln Leu Gln
420
430 420 425 430 Lys Glu Pro Pro Ala Gly Ser Cys Met Leu Gly Gly Leu Ser Leu Gln
435 440 445 440 445 Glu Val Thr Ser Leu Ala Met Glu Glu Ser Gln Glu Ala Lys Ser Leu 450 455 460 His Gln Pro Leu Gly Ile Cys Thr Asp Arg Thr Ser Asp Pro Asn Val 470 475 Leu His Ser Gly Glu Glu Gly Thr Pro Gln Tyr Leu Lys Gly Gln Leu 485 490 Pro Leu Leu Ser Ser Val Gln Ile Glu Gly His Pro Met Ser Leu Pro 505 Leu Gln Pro Pro Ser Gly Pro Cys Ser Pro Ser Asp Gln Gly Pro Ser 515 520 525 Pro Trp Gly Leu Leu Glu Ser Leu Val Cys Pro Lys Asp Glu Ala Lys 530 540 Ser Pro Ala Pro Glu Thr Ser Asp Leu Glu Gln Pro Thr Glu Leu Asp

Ser Leu Phe Arg Gly Leu Ala Leu Thr Val Gln Trp Glu Ser 565 570 <210> 3 <211> 211 <212> PRT <213> Homo sapiens <400> 3 Pro Glu Asp Pro Ser Asp Leu Leu Gln His Val Lys Phe Gln Ser Ser Asn Phe Glu Asn Ile Leu Thr Trp Asp Ser Gly Pro Glu Gly Thr Pro Asp Thr Val Tyr Ser Ile Glu Tyr Lys Thr Tyr Gly Glu Arg Asp Trp 35 40 45 35 Val Ala Lys Lys Gly Cys Gln Arg Ile Thr Arg Lys Ser Cys Asn Leu 50 60 Thr Val Glu Thr Gly Asn Leu Thr Glu Leu Tyr Tyr Ala Arg Val Thr 65 70 75 80 Ala Val Ser Ala Gly Gly Arg Ser Ala Thr Lys Met Thr Asp Arg Phe 85 90 95 85 Ser Ser Leu Gln His Thr Thr Leu Lys Pro Pro Asp Val Thr Cys Ile 110ĹŨŨ 105 Ser Lys Val Arg Ser Ile Gln Met Ile Val His Pro Thr Pro 115 120 125 Ile Arg Ala Gly Asp Gly His Arg Leu Thr Leu Glu Asp Ile Phe His
130 140 Asp Leu Phe Tyr His Leu Glu Leu Gln Val Asn Arg Thr Tyr Gln Met 145 150 160 150 His Leu Gly Gly Lys Gln Arg Glu Tyr Glu Phe Phe Gly Leu Thr Pro 175 165 170 Asp Thr Glu Phe Leu Gly Thr Ile Met Ile Cys Val Pro Thr Trp Ala 190 185 180 Lys Glu Ser Ala Pro Tyr Met Cys Arg Val Lys Thr Leu Pro Asp Arg 195 200 205 Thr Trp Thr 210 <210> 4 <211> 6 <212> PRT <213> Artificial Sequence <220> <223> Glu-Glu peptide tag <400> 4 Glu Tyr Met Pro Met Glu <210> 5 <211> 8 <212> PRT <213> Artificial Sequence <220> <223> Flag-tag peptide <400> 5 Asp Tyr Lys Asp Asp Asp Lys 1 <210> 6 <211> 699

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<213> Homo sapiens

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| gago gggg acco aact taca ggca atct gato gaca ccco aggt | gcaco cctga tggta aacag aagga tccaa gagco gtgco tggca | egt agg acg gca agt agg egg agc | cagte tcace tggae cgtae acaae ccaae ccaae actce aggge | cttco atgco ccgto gtgco agggo gaaco gtggo gaaco gaaco | ot ci gt gg gt gg ga gg ca gg ga ga gg ci gt ci | ttcco gtggf gtca gtcto cccco gtca gtca tcctf | ccca tggad tgcat gcgtd ccaad gagaa gcctg tcttd | a aaa gtg c aaa c caa g aca g caa c ctaa | accca gagco cacco acagco acago ctgco ctaca cgtga | aagg aaga gtcc gtgt gtgga agga atgc | acada acada agci | cccto acccto accao accto accto acctao tcaco | at ga gga gga at gcc tt aa | gatct ggtca ggagg ctggg cgaga cccat gacca ggaca | gccgag tcccgg aagttc gagcag ttgaat aaaacc tcccgg cccagc acgcct aagagc | 120 180 240 300 360 420 480 540 600 |
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| ctt Leu | atg Met | ggg Gly | acc Thr 15 | ctg Leu | gcc Ala | acc Thr | agc Ser | tgc Cys 20 | ctc Leu | ctt Leu | ctc Leu | ttg Leu | gcc Ala 25 | ctc Leu | ttg Leu | 101 |
| | | | | | | | | | | | | | | ctt Leu | | 149 |
| aag Lys | tcc Ser 45 | aac Asn | ttc Phe | cag Gln | cag Gln | ccc Pro 50 | tat Tyr | atc Ile | acc Thr | aac Asn | cgc Arg 55 | acc Thr | ttc Phe | atg Met | ctg Leu | 197 |
| gct Ala 60 | aag Lys | gag Glu | gct Ala | agc Ser | ttg Leu 65 | gct Ala | gat Asp | aac Asn | aac Asn | aca Thr 70 | gac Asp | gtt val | cgt Arg | ctc Leu | att Ile 75 | 245 |
| ggg Gly | gag Glu | aaa Lys | ctg Leu | ttc Phe 80 | His | Gly | gtc Val | Ser | Met | Ser | gag Glu | cgc Arg | tgc Cys | tat Tyr 90 | ctg Leu | 293 |
| | | | | | | | | | | | | | | cct Pro | | 341 |
| tct Ser | gat Asp | agg Arg 110 | ttc Phe | cag Gln | cct Pro | tat Tyr | atg Met 115 | cag Gln | gag Glu | gtg Val | gtg val | ccc Pro 120 | ttc Phe | ctg Leu | gcc Ala | 389 |
| agg Arg | ctc Leu 125 | agc Ser | aac Asn | agg Arg | cta Leu | agc Ser 130 | aca Thr | tgt Cys | cat His | att Ile | gaa Glu 135 | ggt Gly | gat Asp | gac Asp | ctg Leu | 437 |
| | | | | | | | | | | | | | | aag Lys | | 485 |
| gga Gly | gag Glu | agt Ser | gga Gly | gag Glu 160 | atc Ile | aaa Lys | gca Ala | att Ile | gga Gly 165 | gaa Glu | ctg Leu | gat Asp | ttg Leu | ctg Leu 170 | ttt Phe | 533 |
| | | | | | | | | | | | | | | | | |

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His His His His His
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| gca ttg gga atg gta cca cct ccc gaa aat gtc aga atg aat tct gtt Ala Leu Gly Met Val Pro Pro Pro Glu Asn Val Arg Met Asn Ser Val 20 25 30 | 96 | | | | | | | | | | | | | |
| aat ttc aag aac att cta cag tgg gag tca cct gct ttt gcc aaa ggg Asn Phe Lys Asn Ile Leu Gln Trp Glu Ser Pro Ala Phe Ala Lys Gly 35 40 45 | 144 | | | | | | | | | | | | | |
| aac ctg act ttc aca gct cag tac cta agt tat agg ata ttc caa gat Asn Leu Thr Phe Thr Ala Gln Tyr Leu Ser Tyr Arg Ile Phe Gln Asp 50 55 60 | 192 | | | | | | | | | | | | | |
| aaa tgc atg aat act acc ttg acg gaa tgt gat ttc tca agt ctt tcc Lys Cys Met Asn Thr Thr Leu Thr Glu Cys Asp Phe Ser Ser Leu Ser 65 70 75 80 | 240 | | | | | | | | | | | | | |
| aag tat ggt gac cac acc ttg aga gtc agg gct gaa ttt gca gat gag Lys Tyr Gly Asp His Thr Leu Arg Val Arg Ala Glu Phe Ala Asp Glu 85 90 95 | 288 | | | | | | | | | | | | | |
| cat tca gac tgg gta aac atc acc ttc tgt cct gtg gat gac acc att His Ser Asp Trp Val Asn Ile Thr Phe Cys Pro Val Asp Asp Thr Ile 100 105 110 | 336 | | | | | | | | | | | | | |
| att gga ccc cct gga atg caa gta gaa gta ctt gat gat tct tta cat Ile Gly Pro Pro Gly Met Gln Val Glu Val Leu Asp Asp Ser Leu His 115 120 125 | 384 | | | | | | | | | | | | | |
| atg cgt ttc tta gcc cct aaa att gag aat gaa tac gaa act tgg act Met Arg Phe Leu Ala Pro Lys Ile Glu Asn Glu Tyr Glu Thr Trp Thr 130 135 140 | 432 | | | | | | | | | | | | | |
| atg aag aat gtg tat aac tca tgg act tat aat gtg caa tac tgg aaa Met Lys Asn Val Tyr Asn Ser Trp Thr Tyr Asn Val Gln Tyr Trp Lys 145 150 155 160 | 480 | | | | | | | | | | | | | |
| aac ggt act gat gaa aag ttt caa att act ccc cag tat gac ttt gag Asn Gly Thr Asp Glu Lys Phe Gln Ile Thr Pro Gln Tyr Asp Phe Glu 165 170 175 | 528 | | | | | | | | | | | | | |
| gtc ctc aga aac ctg gag cca tgg aca act tat tgt gtt caa gtt cga Val Leu Arg Asn Leu Glu Pro Trp Thr Thr Tyr Cys Val Gln Val Arg 180 185 190 | 576 | | | | | | | | | | | | | |

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ggg ttt ctt cct gat cgg aac aaa gct ggg gaa tgg agt gag cct gtc
Gly Phe Leu Pro Asp Arg Asn Lys Ala Gly Glu Trp Ser Glu Pro Val
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tgt gag caa aca acc cat gac gaa acg gtc ccc tcc
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Ala Leu Gly Met Val Pro Pro Pro Glu Asn Val Arg Met Asn Ser Val
20 25 30
Asn Phe Lys Asn Ile Leu Gln Trp Glu Ser Pro Ala Phe Ala Lys Gly
                                 40
Asn Leu Thr Phe Thr Ala Gln Tyr Leu Ser Tyr Arg Ile Phe Gln Asp 50 60
Lys Cys Met Asn Thr Thr Leu Thr Glu Cys Asp Phe Ser Ser Leu Ser 65 70 75 80
Lys Tyr Gly Asp His Thr Leu Arg Val Arg Ala Glu Phe Ala Asp Glu
85 90 95
His Ser Asp Trp Val Asn Ile Thr Phe Cys Pro Val Asp Asp Thr Ile
100 105 110
Ile Gly Pro Pro Gly Met Gln Val Glu Val Leu Asp Asp Ser Leu His
115 120 125
                                 120
Met Arg Phe Leu Ala Pro Lys Ile Glu Asn Glu Tyr Glu Thr Trp Thr
130 135 140
Met Lys Asn Val Tyr Asn Ser Trp Thr Tyr Asn Val Gln Tyr Trp Lys
145 150 155 160
Ash Gly Thr Asp Glu Lys Phe Gln Ile Thr Pro Gln Tyr Asp Phe Glu
                                           170
                   165
Val Leu Arg Asn Leu Glu Pro Trp Thr Thr Tyr Cys Val Gln Val Arg
180 185 190
Gly Phe Leu Pro Asp Arg Asn Lys Ala Gly Glu Trp Ser Glu Pro Val
Cys Glu Gln Thr Thr His Asp Glu Thr Val Pro Ser
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| <220> <221> CDS <222> (1)(1428) | | | | | | | | | | | | | | |
| <223> CRF2-4 extracellular cytokine binding domain fused to IgGg1 with a 6-HIS tag | | | | | | | | | | | | | | |
| <400> 22 atg gcg tgg a Met Ala Trp S 1 | agt ctt ggg Ser Leu Gly 5 | agc tgg ctg Ser Trp Lei | ggt ggc Gly Gly 10 | tgc ctg ctg Cys Leu Leu | gtg tca 48 Val Ser 15 | | | | | | | | | |
| gca ttg gga a Ala Leu Gly M | | | ASN Val | | | | | | | | | | | |
| aat ttc aag a Asn Phe Lys A 35 | | | | | | 4 | | | | | | | | |
| aac ctg act t Asn Leu Thr F 50 | | | | | | 2 | | | | | | | | |
| aaa tgc atg a Lys Cys Met A 65 | aat act acc Asn Thr Thr 70 | ttg acg gaa Leu Thr Glu | a tgt gat u Cys Asp 75 | ttc tca agt Phe Ser Ser | ctt tcc 24 Leu Ser 80 | 0 | | | | | | | | |
| aag tat ggt g Lys Tyr Gly A | | | | | | 8 | | | | | | | | |
| cat tca gac t His Ser Asp T 1 | tgg gta aac Trp Val Asn 100 | atc acc tto Ile Thr Pho 10 | Cys Pro | gtg gat gac Val Asp Asp 110 | acc att 33 Thr Ile | 6 | | | | | | | | |
| att gga ccc o Ile Gly Pro F 115 | cct gga atg Pro Gly Met | caa gta gaa Gln Val Glu 120 | a gta ctt u Val Leu | gat gat tct Asp Asp Ser 125 | tta cat 38 Leu His | 4 | | | | | | | | |
| atg cgt ttc t Met Arg Phe L 130 | Leu Ăla Pro | | ú Asn Glu | | | 2 | | | | | | | | |
| atg aag aat g Met Lys Asn V 145 | gtg tat aac Val Tyr Asn 150 | tca tgg act Ser Trp Thi | t tat aat Tyr Asn 155 | gtg caa tac Val Gln Tyr | tgg aaa 48 Trp Lys 160 | 0 | | | | | | | | |
| aac ggt act g Asn Gly Thr A | gat gaa aag Asp Glu Lys 165 | ttt caa att Phe Gln Ile | t act ccc Thr Pro 170 | cag tat gac Gln Tyr Asp | ttt gag 52 Phe Glu 175 | 8 | | | | | | | | |
| gtc ctc aga a Val Leu Arg A | | | r Thr Tyr | | | 6 | | | | | | | | |
| ggg ttt ctt c Gly Phe Leu F 195 | cct gat cgg Pro Asp Arg | aac aaa gc Asn Lys Ala 200 | t ggg gaa a Gly Glu | tgg agt gag Trp Ser Glu 205 | cct gtc 62 Pro Val | 4 | | | | | | | | |
| tgt gag caa a Cys Glu Gln 1 210 | aca acc cat Thr Thr His | gac gaa acq Asp Glu Thi 215 | g gtc ccc r val Pro | tcc gga tcc Ser Gly Ser 220 | ggt tcg 67 Gly Ser | 2 | | | | | | | | |

| | | | | ٠ | | | | | | 12 | | | | | | |
|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|--------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|------|
| ggt Gly 225 | tcg Ser | ggt Gly | tcg Ser | gag Glu | ccc Pro 230 | aga Arg | tca Ser | tca Ser | gac Asp | aaa | act Thr | cac His | aca Thr | tgc Cys | cca Pro 240 | 720 |
| ccg Pro | tgc Cys | cca Pro | gca Ala | cct Pro 245 | gaa Glu | gcc Ala | gag Glu | ggg Gly | gca Ala 250 | ccg Pro | tca Ser | gtc Val | ttc Phe | ctc Leu 255 | ttc Phe | 768 |
| ccc Pro | cca Pro | aaa Lys | ccc Pro 260 | aag Lys | gac Asp | acc Thr | ctc Leu | atg Met 265 | atc Ile | tcc Ser | cgg Arg | acc Thr | cct Pro 270 | gag Glu | gtc val | 816 |
| aca Thr | tgc Cys | gtg Val 275 | gtg Val | gtg Val | gac Asp | gtg Val | agc Ser 280 | cac His | gaa Glu | gac Asp | cct Pro | gag Glu 285 | gtc Val | aag Lys | ttc Phe | 864 |
| aac Asn | tgg Trp 290 | tac Tyr | gtg Val | gac Asp | ggc Gly | gtg Val 295 | gag Glu | gtg Val | cat His | aat Asn | gcc Ala 300 | aag Lys | aca Thr | aag Lys | ccg Pro | 912 |
| cgg Arg 305 | gag Glu | gag Glu | cag Gln | tac Tyr | aac Asn 310 | agc Ser | acg Thr | tac Tyr | cgt Arg | gtg Val 315 | gtc Val | agc Ser | gtc val | ctc Leu | acc Thr 320 | 960 |
| | | | | | | | aat Asn | | | | | | | | | 1008 |
| | | | | | | | tcc Ser | | | | | | | | | 1056 |
| aaa Lys | ggg Gly | cag Gln 355 | ccc Pro | cga Arg | gaa Glu | cca Pro | cag Gln 360 | gtg val | tac Tyr | acc Thr | ctg Leu | ccc Pro 365 | cca Pro | tcc Ser | cgg Arg | 1104 |
| gat Asp | gag Glu 370 | ctg Leu | acc Thr | aag Lys | aac Asn | cag Glr: 375 | gtc Val | agc Ser | ctg Leu | acc The | tgc cyc 580 | ctg Leu | gtc Val | aaa Lys | ggc Gly | 1152 |
| ttc Phe 385 | tat Tyr | ccc Pro | agc Ser | gac Asp | atc Ile 390 | gcc Ala | gtg Val | gag Glu | tgg Trp | gag Glu 395 | agc Ser | aat Asn | ggg Gly | cag Gln | ccg Pro 400 | 1200 |
| | | | | | | | cct Pro | | | | | | | | | 1248 |
| ttc Phe | ttc Phe | ctc Leu | tac Tyr 420 | agc Ser | aag Lys | ctc Leu | acc Thr | gtg Val 425 | gac Asp | aag Lys | agc Ser | agg Arg | tgg Trp 430 | cag Gln | cag Gln | 1296 |
| ggg Gly | aac Asn | gtc Val 435 | ttc Phe | tca Ser | tgc Cys | tcc Ser | gtg Val 440 | atg Met | cat His | gag Glu | gct Ala | ctg Leu 445 | cac His | aac Asn | cac His | 1344 |
| | | | | | | | ctg Leu | | | | | | | | | 1392 |
| | | | | | | | cac His | | | | | | | | | 1428 |
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<210> 23 <211> 476 <212> PRT <213> Homo sapiens

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35 40 45 Asn Leu Thr Phe Thr Ala Gln Tyr Leu Ser Tyr Arg Ile Phe Gln Asp 50 60 Lys Cys Met Asn Thr Thr Leu Thr Glu Cys Asp Phe Ser Ser Leu Ser 65 70 75 80 Lys Tyr Gly Asp His Thr Leu Arg Val Arg Ála Glu Phe Ala Asp Glu 85_ 90 95 His Ser Asp Trp Val Asn Ile Thr Phe Cys Pro Val Asp Asp Thr Ile 100 105 110 ... Ile Gly Pro Pro Gly Met Gln Val Glu Val Leu Asp Asp Ser Leu His 115 120 125 120 Met Arg Phe Leu Ala Pro Lys Île Glu Asn Glu Tyr Glu Thr Trp Thr 130 135 140 Met Lys Asn Val Tyr Asn Ser Trp Thr Tyr Asn Val Gln Tyr Trp Lys 145 150 160 Asn Gly Thr Asp Glu Lys Phe Gln Ile Thr Pro Gln Tyr Asp Phe Glu 165 170 175 Val Leu Arg Asn Leu Glu Pro Trp Thr Thr Tyr Cys Val Gln Val Arg Gly Phe Leu Pro Asp Arg Asn Lys Ala Gly Glu Trp Ser Glu Pro Val Cys Glu Gln Thr Thr His Asp Glu Thr Val Pro Ser Gly Ser Gly Ser 210 220 220 Gly Ser Gly Ser Glu Pro Arg Ser Ser Asp Lys Thr His Thr Cys Pro 240 Pro Cys Pro Ala Pro Glu Ala Glu Gly Ala Pro Ser Val Phe Leu Phe 245 250 250 Pro Pro Lys Pro Lys Asp Thr Leu Met Ile Ser Arg Thr Pro Glu Val 260 Thr Cys Val Val Val Asp Val Ser His Glu Asp Pro Glu Val Lys Phe 275 280 285 Asn Trp Tyr Val Asp Gly Val Glu Val His Asn Ala Lys Thr Lys Pro 290 295 300 Arg Glu Glu Gln Tyr Asn Ser Thr Tyr Arg Val Val Ser Val Leu Thr 305 310 320 Val Leu His Gln Asp Trp Leu Asn Gly Lys Glu Tyr Lys Cys Lys Val 325 330 335 Ser Asn Lys Ala Leu Pro Ser Ser Ile Glu Lys Thr Ile Ser Lys Ala 345 Lys Gly Gln Pro Arg Glu Pro Gln Val Tyr Thr Leu Pro Pro Ser Arg 355 360 365 Asp Glu Leu Thr Lys Asn Gln Val Ser Leu Thr Cys Leu Val Lys Gly 370 375 380 Phe Tyr Pro Ser Asp Ile Ala Val Glu Trp Glu Ser Asn Gly Gln Pro 385 395 Glu Asn Asn Tyr Lys Thr Thr Pro Pro Val Leu Asp Ser Asp Gly Ser 405 410 415 Phe Phe Leu Tyr Ser Lys Leu Thr Val Asp Lys Ser Arg Trp Gln Gln 420 430 Gly Asn Val Phe Ser Cys Ser Val Met His Glu Ala Leu His Asn His 435 445 Tyr Thr Gln Lys Ser Leu Ser Leu Ser Pro Gly Lys Leu Val Pro Arg 450 460 435 Gly ser Gly Ser Gly Gly His His His His His

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14

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      fused to IgGg1 with a Glu-Glu tag
<221> CDS
<222> (1)...(1452)
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96 gcc cct gag gac ccc tcg gat ctg ctc cag cac gtg aaa ttc cag tcc Ăla Pro Ğlü Ăsp Pro Ser Āsp Leŭ Leu Glñ His Val Lys Phe Glň Ser agc aac ttt gaa aac atc ctg acg tgg gac agc ggg cca gag ggc acc Ser Asn Phe Glu Asn Ile Leu Thr Trp Asp Ser Gly Pro Glu Gly Thr 144 40 cca gac acg gtc tac agc atc gag tat aag acg tac gga gag gac Pro Asp Thr Val Tyr Ser Ile Glu Tyr Lys Thr Tyr Gly Glu Arg Asp 192 tgg gtg gca aag aag ggc tgt cag cgg atc acc cgg aag tcc tgc aac Trp Val Ala Lys Lys Gly Cys Gln Arg Ile Thr Arg Lys Ser Cys Asn 65 70 75 80 240 ctg acg gtg gag acg ggc aac ctc acg gag ctc tac tat gcc agg gtc Leu Thr Val Glu Thr Gly Asn Leu Thr Glu Leu Tyr Tyr Ala Arg Val 288 acc gct gtc agt gcg gga ggc cgg tca gcc acc aag atg act gac agg Thr Ala Val Ser Ala Gly Gly Arg Ser Ala Thr Lys Met Thr Asp Arg 336 384 ttc agc tct ctg cag cac act acc ctc aag cca cct gat gtg acc tgt Phe Ser Ser Leu Gln His Thr Thr Leu Lys Pro Pro Asp Val Thr Cys atc tcc aaa gtg aga tcg att cag atg att gtt cat cct acc ccc acg Ile Ser Lys Val Arg Ser Ile Gln Met Ile Val His Pro Thr Pro Thr 432 cca atc cgt gca ggc gat ggc cac cgg cta acc ctg gaa gac atc ttc Pro Ile Arg Ala Gly Asp Gly His Arg Leu Thr Leu Glu Asp Ile Phe 480 cat gac ctg ttc tac cac tta gag ctc cag gtc aac cgc acc tac caa His Asp Leu Phe Tyr His Leu Glu Leu Gln Val Asn Arg Thr Tyr Gln 528 170 atg cac ctt gga ggg aag cag aga gaa tat gag ttc ttc ggc ctg acc Met His Leu Gly Gly Lys Gln Arg Glu Tyr Glu Phe Phe Gly Leu Thr 576 cct gac aca gag ttc ctt ggc acc atc atg att tgc gtt ccc acc tgg Pro Asp Thr Glu Phe Leu Gly Thr Ile Met Ile Cys Val Pro Thr Trp 624 gcc aag gag agt gcc ccc tac atg tgc cga gtg aag aca ctg cca gac Ala Lys Glu Ser Ala Pro Tyr Met Cys Arg Val Lys Thr Leu Pro Asp 210 220 cgg aca tgg acc gga tcc ggt tcg ggt tcg ggt tcg gag ccc aga tca Arg Thr Trp Thr Gly Ser Gly Ser Gly Ser Glu Pro Arg Ser 225 230 235 240 720 tca gac aaa act cac aca tgc cca ccg tgc cca gca cct gaa gcc gag Ser Asp Lys Thr His Thr Cys Pro Pro Cys Pro Ala Pro Glu Ala Glu 245 250 255 768 ggg gca ccg tca gtc ttc ctc ttc ccc cca aaa ccc aag gac acc ctc Gly Ala Pro Ser Val Phe Leu Phe Pro Pro Lys Pro Lys Asp Thr Leu 260 265 270816 atg atc tcc cgg acc cct gag gtc aca tgc gtg gtg gtg gac gtg agc Met Ile Ser Arg Thr Pro Glu Val Thr Cys Val Val Val Asp Val Ser 275 280 285 864

| | | | | gag Glu | | | | | | | | | | | | 912 |
|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|------|
| gtg val 305 | cat His | aat Asn | gcc Ala | aag Lys | aca Thr 310 | aag Lys | ccg Pro | cgg Arg | gag Glu | gag Glu 315 | cag Gln | tac Tyr | aac Asn | agc Ser | acg Thr 320 | 960 |
| | | | | agc Ser 325 | | | | | | | | | | | | 1008 |
| ggc Gly | aag Lys | gag Glu | tac Tyr 340 | aag Lys | tgc Cys | aag Lys | gtc Val | tcc Ser 345 | aac Asn | aaa Lys | gcc Ala | ctc Leu | cca Pro 350 | tcc Ser | tcc Ser | 1056 |
| atc Ile | gag Glu | aaa Lys 355 | acc Thr | atc Ile | tcc Ser | aaa Lys | gcc Ala 360 | aaa Lys | ggg Gly | cag Gln | ccc Pro | cga Arg 365 | gaa Glu | cca Pro | cag Gln | 1104 |
| gtg val | tac Tyr 370 | acc Thr | ctg Leu | ccc Pro | cca Pro | tcc ser 375 | cgg Arg | gat Asp | gag Glu | ctg Leu | acc Thr 380 | aag Lys | aac Asn | cag Gln | gtc Val | 1152 |
| | | | | ctg Leu | | | | | | | | | | | | 1200 |
| gag Glu | tgg Trp | gag Glu | agc Ser | aat Asn 405 | ggg Gly | cag Gln | ccg Pro | gag Glu | aac Asn 410 | aac Asn | tac Tyr | aag Lys | acc Thr | acg Thr 415 | cct Pro | 1248 |
| ccc Pro | gtg val | ctg Leu | gac Asp 420 | tcc Ser | gac Asp | ggc Gly | tcc Ser | ttc Phe 425 | ttc Phe | ctc Leu | tac Tyr | agc Ser | aag Lys 430 | ctc Leu | acc Thr | 1296 |
| gtg Val | gac Asp | aag Lys 435 | agc Ser | agg Arg | tgg Trp | cag Gln | cag Gln 440 | ggg Gly | aac Asn | gtc Val | ttc Phe | tca Ser 445 | tgc Cys | tcc Ser | gtg Val | 1344 |
| atg Met | cat His 450 | gag Glu | gct Ala | ctg Leu | cac His | aac Asn 455 | cac His | tac Tyr | acg Thr | cag Gln | aag Lys 460 | agc Ser | ctc Leu | tcc Ser | ctg Leu | 1392 |
| tct Ser 465 | ccg Pro | ggt Gly | aaa Lys | ctg Leu | gtt Val 470 | ccg Pro | cgt Arg | ggt Gly | tcc Ser | gga Gly 475 | tca Ser | ggt Gly | ggc Gly | gag Glu | tat Tyr 480 | 1440 |
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<212> PRT <213> Artificial Sequence

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<213> Artificial Sequence

<223> Oligonucleotide primer ZC37693

<400> 31

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                                                                          23
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20 25 30
Phe Thr Ala Gin Tyr Leu Ser Tyr Arg Ile Phe Gin Asp Lys Cys Met 35
Asn Thr Thr Leu Thr Glu Cys Asp Phe Ser Ser Leu Ser Lys Tyr Gly 50 60
Asp His Thr Leu Arg Val Arg Ala Glu Phe Ala Asp Glu His Ser Asp 65 70 75 80
Trp Val Asn Ile Thr Phe Cys Pro Val Asp Asp Thr Ile Ile Gly Pro 85 90 95
Pro Gly Met Gln Val Glu Val Leu Ala Asp Ser Leu His Met Arg Phe 100 105 110
Leu Ala Pro Lys Ile Glu Asn Glu Tyr Glu Thr Trp Thr Met Lys Asn 115
Val Tyr Asn Ser Trp Thr Tyr Asn Val Gln Tyr Trp Lys Asn Gly Thr
130 135 140
Asp Glu Lys Phe Gln Ile Thr Pro Gln Tyr Asp Phe Glu Val Leu Arg
145 150 155 160
Asn Leu Glu Pro Trp Thr Thr Tyr Cys Val Gln Val Arg Gly Phe Leu
165 170 175
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Pro Asp Arg Asn Lys Ala Gly Glu Trp Ser Glu Pro Val Cys Glu Gln
Thr Thr His Asp Glu Thr Val
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1 10 15
Glu Ala Glu Phe Phe His His Ile Leu His Trp Thr Pro Ile Pro Asn
20 25 30
Gln Ser Glu Ser Thr Cys Tyr Glu Val Ala Leu Leu Arg Tyr Gly Ile
35 40 45
Glu Ser Trp Asn Ser Ile Ser Asn Cys Ser Gln Thr Leu Ser Tyr Asp 50 60
Leu Thr Ala Val Thr Leu Asp Leu Tyr His Ser Asn Gly Tyr Arg Ala 65 70 75 80
Arg Val Arg Ala Val Asp Gly Ser Arg His Ser Asn Trp Thr Val Thr
                                        90
                                                              95
                 85
Asn Thr Arg Phe Ser Val Asp Glu Val Thr Leu Thr Val Gly Ser Val 100 105 110
Asn Leu Glu Ile His Asn Gly Phe Ile Leu Gly Lys Ile Gln Leu Pro
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19 Arg Pro Lys Met Ala Pro Ala Ash Asp Thr Tyr Glu Ser Ile Phe Ser 130 135 140 His Phe Arg Glu Tyr Glu Île Ala Île Arg Lys Val Pro Gly Asn Phe 155 150 Thr Phe Thr His Lys Lys Val Lys His Glu Asn Phe Ser Leu Leu Thr 165 170 175 Ser Gly Glu Val Gly Glu Phe Cys Val Gln Val Lys Pro Ser Val Ala 180 185 190 190 180 185 Ser Arg Ser Asn Lys Gly Met Trp Ser Lys Glu Glu Cys Ile Ser Leu 195 200 205 Thr Arg Gln 21Ō

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